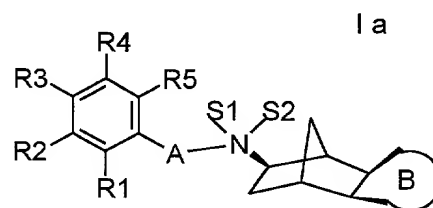
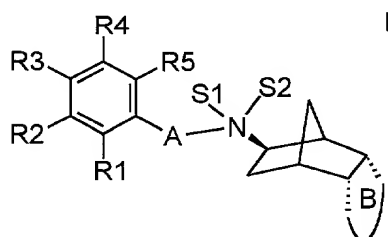


What is claimed is:

1. A substituted norbornylamino derivative having exo-configured nitrogen and an endo-fused five-, six- or seven-membered ring of the formula I or a pharmaceutically acceptable salt or trifluoroacetate thereof, or having exo-configured nitrogen and an exo-fused five-, six- or seven-membered ring of the formula I a or a pharmaceutically acceptable salt or trifluoroacetate thereof



in which:

A is (C<sub>1</sub>-C<sub>4</sub>)-alkylene;

S1 is a free electron pair or (C<sub>1</sub>-C<sub>4</sub>)-alkyl;

S2 is (C<sub>1</sub>-C<sub>4</sub>)-alkyl or H;

where, if S1 and S2 are alkyl, X<sup>-</sup> in the resulting grouping [N<sup>+</sup>(S1S2)-X<sup>-</sup>] corresponds to a pharmacologically acceptable anion or trifluoroacetate;

B is a saturated or unsaturated five-, six- or seven-membered carbon ring which may be mono- or, independently of one another, polysubstituted by oxo, hydroxyl, (C<sub>1</sub>-C<sub>4</sub>)-alkoxy and (C<sub>1</sub>-C<sub>4</sub>)-alkyl;

and

R1, R2, R3, R4 and R5

are, independently of one another, H, OH, F, Cl, Br, I, CN, NO<sub>2</sub>, amidino, -CO<sub>2</sub>R(11), -CONR(11)R(12), -SO<sub>r</sub>R(11), -SO<sub>s</sub>NR(11)-R(12), (C<sub>1</sub>-C<sub>4</sub>)-alkyl, (C<sub>1</sub>-C<sub>4</sub>)-alkoxy, (C<sub>1</sub>-C<sub>4</sub>)-alkoxy-(C<sub>1</sub>-C<sub>4</sub>)-alkyl, (C<sub>1</sub>-C<sub>4</sub>)-alkoxy-(C<sub>1</sub>-C<sub>4</sub>)-alkyloxy, hydroxy-(C<sub>1</sub>-C<sub>4</sub>)-alkyl, (C<sub>3</sub>-C<sub>7</sub>)-cycloalkoxy or phenyloxy,

where phenyl is unsubstituted or substituted by up to three substituents, which are independent of one another and are F, Cl, Br, or methoxy;

amino, (C<sub>1</sub>-C<sub>4</sub>)-alkylamino, di-(C<sub>1</sub>-C<sub>4</sub>)-alkylamino, amino-(C<sub>1</sub>-C<sub>4</sub>)-alkyl, di-(C<sub>1</sub>-C<sub>4</sub>)-alkylamino-(C<sub>1</sub>-C<sub>4</sub>)-alkyl, (C<sub>1</sub>-C<sub>4</sub>)-alkylamino-(C<sub>1</sub>-C<sub>4</sub>)-alkyl,

where some or all of the hydrogen atoms in the alkyl radicals may be substituted by fluorine;

R11 and R12

are, independently of one another, H or (C<sub>1</sub>-C<sub>4</sub>)-alkyl,

where some or all of the hydrogen atoms in the alkyl radicals may be substituted by fluorine;

r is 0, 1 or 2;

s is 1 or 2;

or

R1 and R2, R2 and R3, R3 and R4 or R4 and R5

in each case together are a group -O-(CH<sub>2</sub>)<sub>n</sub>-O-;

n is 1 or 2;

and

the radicals R1, R2, R3, R4 or R5 which remain in each case

are, independently of one another, H, OH, F, Cl, Br, I, CN, NO<sub>2</sub>,

amidino, -CO<sub>2</sub>R(11), -CONR(11)R(12), -SO<sub>r</sub>R(11), -SO<sub>s</sub>NR(11)-

R(12), (C<sub>1</sub>-C<sub>4</sub>)-alkyl, (C<sub>1</sub>-C<sub>4</sub>)-alkoxy, (C<sub>1</sub>-C<sub>4</sub>)-alkoxy-(C<sub>1</sub>-C<sub>4</sub>)-alkyl,

(C<sub>3</sub>-C<sub>7</sub>)-cycloalkoxy, hydroxy-(C<sub>1</sub>-C<sub>4</sub>)-alkyl, amino, (C<sub>1</sub>-C<sub>4</sub>)-alkyl-

amino, di-(C<sub>1</sub>-C<sub>4</sub>)-alkylamino, amino-(C<sub>1</sub>-C<sub>4</sub>)-alkyl, di-(C<sub>1</sub>-C<sub>4</sub>)-

alkylamino-(C<sub>1</sub>-C<sub>4</sub>)-alkyl, (C<sub>1</sub>-C<sub>4</sub>)-alkylamino-(C<sub>1</sub>-C<sub>4</sub>)-alkyl,

where some or all of the hydrogen atoms in the alkyl

radicals may be substituted by fluorine;

R11 and R12

are, independently of one another, H or (C<sub>1</sub>-C<sub>4</sub>)-alkyl,

where some or all of the hydrogen atoms in the

alkyl radicals may be substituted by fluorine;

r is 0, 1 or 2;

s is 1 or 2;

except for benzyl(octahydro-4,7-methanoinden-5-yl)amine.

2. A compound of Claim 1, having exo-configured nitrogen and an endo-

fused five- or six-membered ring of the formula I, or having exo-configured

nitrogen and an exo-fused five- or six-membered ring of the formula I a, in

which:

A is (C<sub>1</sub>-C<sub>2</sub>)-alkylene;

S1 is a free electron pair or methyl;

S2 is H;

B is a saturated or unsaturated five- or six-membered carbon ring;

R1, R2, R3, R4 and R5

5 are, independently of one another, H, amino, hydroxymethyl, OH, methoxy, F, Cl, Br or iodine;

or

R2 and R3

together are -O-CH<sub>2</sub>-O-;

10 and

the remaining radicals R1, R4 and R5

are, independently of one another, H, OH, F, Cl, Br, I, CN, NO<sub>2</sub>, (C<sub>1</sub>-C<sub>2</sub>)-alkoxy, amino, (C<sub>1</sub>-C<sub>2</sub>)-alkylamino or di-(C<sub>1</sub>-C<sub>2</sub>)-alkylamino,

15 where some or all of the hydrogen atoms in the alkyl radicals may be substituted by fluorine;

or a pharmaceutically acceptable salt or trifluoroacetate thereof.

3. A compound of Claim 1, having exo-configured nitrogen and an endo-fused five- or six-membered ring of the formula I, or having exo-configured nitrogen and an exo-fused five- or six-membered ring of the formula I a, in which:

A is (C<sub>1</sub>-C<sub>2</sub>)-alkylene;

S1 is a free electron pair;

25 S2 is H;

B is a saturated or unsaturated five- or six-membered carbon ring;

R1, R3 and R5

are hydrogen;

and R2 and R4

30 are, independently of one another, H, methoxy, F or Cl;

or

R2 and R3

together are -O-CH<sub>2</sub>-O-;

and

35 R1, R4 and R5

are hydrogen;

or a pharmaceutically acceptable salt thereof.

4. A compound of Claim 1, having *exo*-configured nitrogen and an *endo*-fused five- or six-membered ring of the formula I, or having *exo*-configured nitrogen and an *exo*-fused five-membered ring of the formula I a, wherein the compound is:

- 5    *exo/endo*-(3-chlorobenzyl)(octahydro-4,7-methanoinden-5-yl)amine,  
      *exo/endo*-(3-fluorobenzyl)(octahydro-4,7-methanoinden-5-yl)amine,  
      *exo/endo*-benzo[1,3]dioxol-5-ylmethyl(octahydro-4,7-methanoinden-5-yl)-  
      amine,  
      *exo/endo*-(*rac*)-(3-methoxybenzyl)(octahydro-4,7-methanoinden-5-yl)-  
10    amine,  
      *exo/endo*-(+)-(3-methoxybenzyl)(octahydro-4,7-methanoinden-5-yl)amine,  
      *exo/endo*-(-)-(3-methoxybenzyl)(octahydro-4,7-methanoinden-5-yl)amine,  
      *exo/endo*-[1-(3-methoxyphenyl)ethyl](octahydro-4,7-methanoinden-5-yl)-  
      amine,  
15    *exo/endo*-(3-fluorobenzyl)(3a,4,5,6,7,7a-hexahydro-1H-4,7-methanoinden-  
      5-yl)amine,  
      *exo/endo*-(3-fluorobenzyl)(3a,4,5,6,7,7a-hexahydro-3H-4,7-methanoinden-  
      5-yl)amine,  
      *exo/endo*-(3a,4,5,6,7,7a-hexahydro-1H-4,7-methanoinden-5-yl)(3-  
20    methoxybenzyl)amine,  
      *exo/endo*-(3a,4,5,6,7,7a-hexahydro-3H-4,7-methanoinden-5-yl)(3-  
      methoxybenzyl)amine,  
      *exo/endo*-(decahydro-1,4-methanonaphthalen-2-yl)(3-methoxybenzyl)-  
      amine,  
25    *exo/endo*-(3,5-difluorobenzyl)(octahydro-4,7-methanoinden-5-yl)amine,  
      *exo/exo*-(3-fluorobenzyl)(octahydro-4,7-methanoinden-5-yl)amine, or  
      *exo/exo*-(3-methoxybenzyl)(octahydro-4,7-methanoinden-5-yl)amine, or  
      a pharmaceutically acceptable salt or trifluoroacetate thereof.
- 30    5. A compound of Claim 1, having *exo*-configured nitrogen and an *endo*-  
      fused 5- or 6-membered ring, wherein the compound is:  
      *exo/endo*-(3-chlorobenzyl)(octahydro-4,7-methanoinden-5-yl)amine,  
      *exo/endo*-(3-fluorobenzyl)(octahydro-4,7-methanoinden-5-yl)amine,  
      *exo/endo*-(3-fluorobenzyl)(3a,4,5,6,7,7a-hexahydro-1H-4,7-methanoinden-  
35    5-yl)amine,  
      *exo/endo*-(3-fluorobenzyl)(3a,4,5,6,7,7a-hexahydro-3H-4,7-methanoinden-  
      5-yl)amine,

exo/endo-benzo[1,3]dioxol-5-ylmethyl(octahydro-4,7-methanoinden-5-yl)-  
amine,

exo/endo-(*rac*)-(3-methoxybenzyl)(octahydro-4,7-methanoinden-5-yl)  
amine,

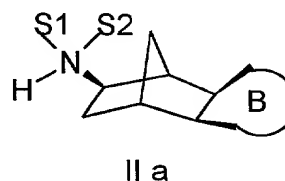
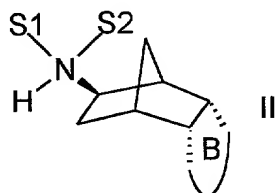
- 5    exo/endo-(+)-(3-methoxybenzyl)(octahydro-4,7-methanoinden-5-yl)amine,  
exo/endo-(decahydro-1,4-methanonaphthalen-2-yl)(3-methoxybenzyl)-  
amine,

exo/endo-(-)-(3-methoxybenzyl)(octahydro-4,7-methanoinden-5-yl)amine,  
or

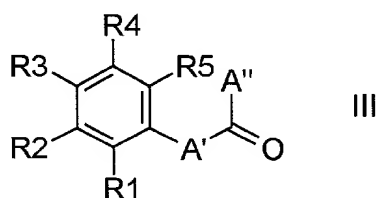
- 10    exo/endo-(3,5-difluorobenzyl)(octahydro-4,7-methanoinden-5-yl)amine, or  
a pharmaceutically acceptable salt or trifluoroacetate thereof.

6. A process for preparing a compound of Claim 1, comprising

- 15    (A) reacting a compound of the formula II or II a



with a compound of the formula III



in which S1, S2, B, R1, R2, R3, R4 and R5 are as defined in Claim  
1, while independently of one another A' is a bond or (C<sub>1</sub>-C<sub>3</sub>)-  
alkylene and A'' is H or (C<sub>1</sub>-C<sub>3</sub>)-alkyl and A' and A'' together with the  
carbon atom of the carbonyl group represent the same number of  
carbon atoms as A,

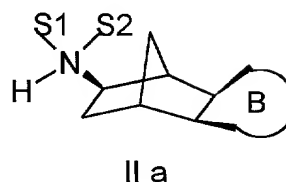
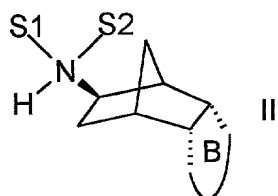
in the presence of suitable reducing agents and optionally also  
Lewis acids directly to give a compound of the formula I or I a, and

(B) optionally converting the compound of formula I or I a into a pharmaceutically acceptable salt or trifluoroacetate.

7. A process for preparing a compound Claim 1, comprising

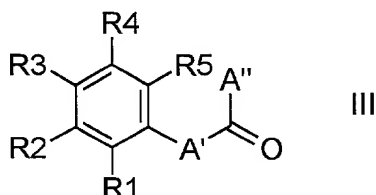
5

(A) reacting a compound of the formula II or II a



10

with a compound of the formula III

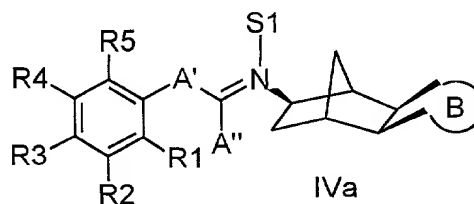
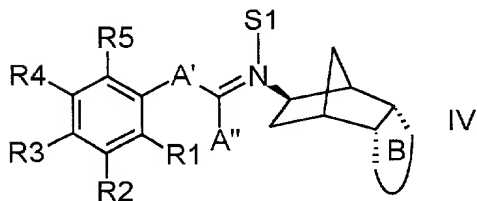


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in which S1, S2, B, R1, R2, R3, R4 and R5 are as defined in Claim 1, while independently of one another A' is a bond or (C<sub>1</sub>-C<sub>3</sub>)-alkylene and A'' is H or (C<sub>1</sub>-C<sub>3</sub>)-alkyl and A' and A'' together with the carbon atom of the carbonyl group represent the same number of carbon atoms as A,

20

(B) isolating the intermediate of the formula IV or IV a



25

formed from the reaction of the compounds of the formulae II or II a and III, in which, if S1 is (C<sub>1</sub>-C<sub>4</sub>)-alkyl, an onium nitrogen is formed which is associated with a counterion,

(C) converting the intermediate with suitable reducing agents into a compound of the formula I or Ia, and

(D) optionally converting the compound of the formula I or Ia into a pharmaceutically acceptable salt or trifluoroacetate.

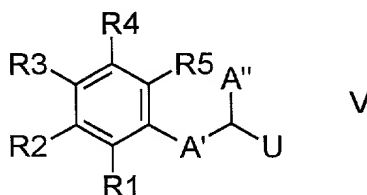
8. A process as claimed in Claim 7, wherein the counterion is chloride or tosylate.

9. A process for preparing a compound of Claim 1, comprising

(A) reacting a compound of the formula II or II a



with an alkylating agent of the formula V



in which U is a nucleophilically substitutable group, and in which S1, S2, B, R1, R2, R3, R4 and R5 are as defined in Claim 1, while independently of one another A' is a bond or (C<sub>1</sub>-C<sub>3</sub>)-alkylene and A'' is H or (C<sub>1</sub>-C<sub>3</sub>)-alkyl and A' and A'' together with the carbon atom to which U is attached represent the same number of carbon atoms as A, to give a compound of the formula I or Ia, and

(B) optionally converting the compound of the formula I or Ia into a pharmaceutically acceptable salt or trifluoroacetate.

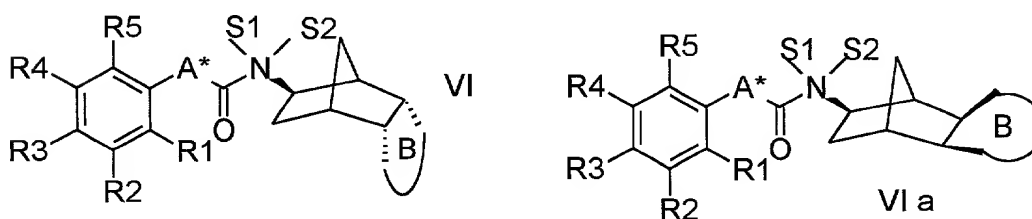
10. A process as claimed in Claim 9, wherein U is chlorine, bromine, iodine, mesylate, tosylate, or triflate.

11. A process as claimed in Claim 9, wherein the reaction step occurs in the presence of one or more non-nucleophilic bases.

12. A process as claimed in Claim 9, wherein the reaction step occurs in the presence of diisopropylethylamine.

13. A process for preparing a compound of Claim 1, comprising

(A) reducing a carboxamide of the formula VI or VI a



in which A\* is a bond or (C<sub>1</sub>-C<sub>3</sub>)-alkylene and the other radicals are as defined in Claim 1 to give a corresponding amine of the formula I or I a, and

(B) optionally converting the amine into a pharmaceutically acceptable salt or trifluoroacetate.

14. A process for converting a secondary amine of the formula I or I a as claimed in claim 1, into a tertiary amine or quaternary ammonium salt, or a pharmaceutically acceptable salt or trifluoroacetate thereof, comprising

(A) mono- or dialkylating a compound of the formula I or Ia in which S1 is a free electron pair and S2 is hydrogen, with alkylating agents of the formula VII



in which S\* is (C<sub>1</sub>-C<sub>4</sub>)-alkyl and U is a nucleophilically substitutable group, thus obtaining a tertiary amine or a quaternary ammonium salt, and

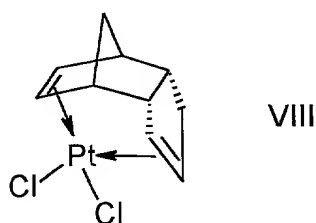


(B) optionally converting the tertiary amine or quaternary ammonium salt into a pharmaceutically acceptable salt or trifluoroacetate.

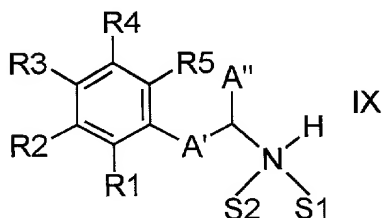
5 15. A process as claimed in Claim 14, wherein U is chlorine, bromine, iodine, mesylate, tosylate, or triflate.

16. A process for preparing a compound Claim 1, comprising

10 (A) reacting a dicyclopentadienylplatinum complex of the formula VIII



with amines of the type of the formula IX



15 in which S1, S2, R1, R2, R3, R4 and R5 are as defined in Claim 1, while independently of one another A' is a bond or (C<sub>1</sub>-C<sub>3</sub>)-alkyl and A'' is H or (C<sub>1</sub>-C<sub>3</sub>)-alkyl and A' and A'' together with the carbon atom to which the nitrogen atom is attached represent the same number of carbon atoms as A, to form an intermediate,

20 (B) reducing the intermediate formed to give a compound of the formula I, and

25 (C) optionally converting the compound into a pharmaceutically acceptable salt or trifluoroacetate.

17. A method of treating or preventing one or more disorders of the respiratory drive, comprising administering an effective amount of a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

5

18. A method as claimed in Claim 17, wherein the disorder is sleep-related.

19. A method as claimed in Claim 18, wherein the disorder is sleep apnea.

10 20. A method of treating or preventing snoring, comprising administering an effective amount of a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

15 21. A method of treating or preventing one or more acute or chronic renal disorders, comprising administering an effective amount of a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

20 22. A method as claimed in Claim 21, wherein the disorder is acute kidney failure, chronic kidney failure, or both.

25 23. A method of treating or preventing impaired intestinal function, comprising administering an effective amount of a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

24. A method of treating or preventing impaired gallbladder function, comprising administering an effective amount of a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

30 25. A method of treating or preventing ischemic states of the peripheral nervous system, comprising administering an effective amount of a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

35 26. A method of treating or preventing ischemic states of the central nervous system, comprising administering an effective amount of a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

27. A method of treating or preventing stroke, comprising administering an effective amount of a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

5

28. A method of treating or preventing ischemic states of peripheral organs and limbs, comprising administering an effective amount of a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

10

29. A method of treating or preventing shock, comprising administering an effective amount of a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

15

30. A method of protecting an organ, comprising protecting the organ with an amount of a compound of formula I or I a as claimed in Claim 1, or a pharmaceutically acceptable salt thereof, effective to reduce or prevent ischemically induced damage to the organ.

20

31. A method of preserving or storing a transplant, comprising placing the transplant into a physiological bath fluid containing a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

25

32. A method of treating or preventing diseases whose primary or secondary cause is cell proliferation, comprising administering an effective amount of a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

30

33. A method of treating or preventing impaired lipid metabolism, comprising administering an effective amount of a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

35

34. A method of treating or preventing infestation by ectoparasites, comprising administering an effective amount of a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

35. A composition comprising a compound of Claim 1 and a pharmaceutically acceptable carrier.

36. A composition comprising a compound of Claim 4 and a pharmaceutically acceptable carrier.

5 37. A composition comprising a compound of Claim 5 and a pharmaceutically acceptable carrier.

38. A method of treating or preventing hypertension, comprising administering an effective amount of a compound of formula I or I a as  
10 claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

39. A method of treating or preventing one or more of the following conditions: coronary vasospasms, atherogenesis and atherosclerosis, left-ventricular hypertrophy and dilated cardiomyopathy, and thrombic  
15 disorders, comprising administering an effective amount of a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

40. A method of preventing biliary calculus, comprising administering an effective amount of a compound of formula I or I a as claimed in Claim 1 or  
20 a pharmaceutically acceptable salt thereof.

41. A method of treating or preventing late diabetic complications, comprising administering an effective amount of a compound of formula I or  
25 I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

42. A method of treating or preventing carcinomatous disorders, comprising administering an effective amount of a compound of formula I or I a as  
30 claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

43. A method of treating or preventing fibrotic disorders, comprising administering an effective amount of a compound of formula I or I a as  
claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

35 44. A method of treating or preventing organ hypertrophies, comprising administering an effective amount of a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.

45. A method of treating or preventing organ hyperplasias, comprising administering an effective amount of a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.
- 5 46. A method of treating or preventing a disease caused by elevated cholesterol levels, comprising administering an effective amount of a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.
- 10 47. A method of treating or preventing a disease caused by endothelial dysfunction, comprising administering an effective amount of a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof.
- 15 48. A method of inhibiting sodium/proton exchanger, subtype 3 (NHE3), in a patient using a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof, comprising giving the patient, for one or more days, up to four doses per day of the compound, wherein the doses are up to 200 mg/kg of body weight.
- 20 49. A method of inhibiting sodium/proton exchanger, subtype 3 (NHE3), in a patient using a compound of formula I or I a as claimed in Claim 1 or a pharmaceutically acceptable salt thereof, comprising giving the patient, for one or more days, a daily dose of the compound of between 0.001 mg/kg and 100 mg/kg of body weight.
- 25 50. A method as claimed in Claim 49, wherein the daily dose is between 1 and 10 mg/kg of body weight.